DOCKET NO.: TIBO-0029 **Application No.:** 09/836,477

Office Action Dated: August 13, 2004

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1. (Currently amended) A method of determining a phenotype of a <u>retrovirus</u>, wherein the retrovirus is the <u>Human Immunodeficiency Virus</u> biological sample, comprising:

- a) obtaining a genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample;
- b) identifying a mutation pattern of the genetic sequence of the <u>Human</u>

 <u>Immunodeficiency Virus</u> biological sample, wherein said mutation pattern comprises at least one mutation that correlates to resistance to at least one therapy;
- c) searching a relational genotype/phenotype database for at least one database mutation pattern similar to said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample;
- d) obtaining at least one database phenotype of the at least one database mutation pattern; and
- e) determining the phenotype of the <u>Human Immunodeficiency Virus</u> biological sample from the at least one database phenotype.
- 2. (Original) The method of claim 1, wherein a series of phenotypes is obtained by repeating steps b) through e) for each therapy in a group of therapies.
- 3. (Currently amended) The method of claim 1, wherein said mutation pattern of the genetic sequence of the Human Immunodeficiency Virus biological sample is specific to a therapy.
- 4. (Currently amended) The method of claim 1, wherein the <u>Human</u>

 <u>Immunodeficiency Virus</u> biological sample is obtained from at least one of a plasma sample, a blood sample, a saliva sample, mucous sample, and a tissue sample.
- 5. (Currently amended) The method of claim 1, wherein the <u>Human</u> <u>Immunodeficiency Virus</u> biological sample comprises at least one of a virus.

6.-7. Canceled

DOCKET NO.: TIBO-0029 **Application No.:** 09/836,477

Office Action Dated: August 13, 2004

8. (Currently amended) The method of claim 1, wherein said at least one mutation is chosen from a frame shift mutation and a generic alteration, a base substitution, and an epigenetic mutation.

9.-12. Canceled

13. (Currently amended) The method of claim [12]1, wherein the genetic sequence of Human Immunodeficiency Virus HIV is chosen from the genetic sequence of the protease region of the Human Immunodeficiency Virus HIV genome, the genetic sequence of the reverse transcriptase region of the Human Immunodeficiency Virus HIV genome, and the genetic sequence of the protease region and reverse transcriptase region of the Human Immunodeficiency Virus HIV genome.

14.-15. Canceled

- 16. (Currently amended) The method of claim 1, wherein said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample comprises at least two mutations that correlate to resistance to at least one therapy.
- 17. (Original) The method of claim 1, wherein the search of the relational genotype/phenotype database for at least one sample with a similar mutation pattern uses cluster searches.
- 18. (Currently amended) The method of claim 1, wherein the database mutation pattern comprises at least one mutation found in said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.
- 19. (Currently amended) The method of claim 1, wherein the database mutation pattern is a mutation pattern in which at least about 50% of the mutations are identical to said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus biological</u> sample.

DOCKET NO.: TIBO-0029 **Application No.:** 09/836,477

Office Action Dated: August 13, 2004

20. (Currently amended) The method of claim 19, wherein the database mutation pattern is a mutation pattern in which at least about 80% of the mutations are identical to said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.

- 21. (Currently amended) The method of claim 20, wherein the database mutation pattern is a mutation pattern in which at least about 90% of the mutations are identical to said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.
- 22. (Currently amended) The method of claim 21, wherein the mutations of the database mutation pattern are identical to said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.
- 23. (Currently amended) The method of claim 1, wherein the phenotype of the Human Immunodeficiency Virus biological sample is a mean fold-change in resistance, wherein said mean fold change is obtained from all of the database phenotypes obtained in step d).
- 24. (Currently amended) The method of claim 1, wherein the phenotype of the Human Immunodeficiency Virus biological sample is expressed as an IC₅₀.
- 25. (Currently amended) A method of reporting a phenotype for a <u>Human</u> <u>Immunodeficiency Virus</u> <u>biological sample</u>, comprising generating a report having the phenotype determined using the method of claim 1.
- 26. (Original) A report, comprising the phenotype determined using the method of claim 1.
- 27. (Original) A computer readable media, comprising the phenotype determined using the method of claim 1.

DOCKET NO.: TIBO-0029 **Application No.:** 09/836,477

Office Action Dated: August 13, 2004

28. (Currently amended) A method of determining a phenotype of a <u>retrovirus</u>, wherein the retrovirus is the <u>Human Immunodeficiency Virus</u> comprising:

- a) obtaining a genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample;
- b) searching a relational genotype/phenotype database for at least one database genetic sequence similar to said genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample;
 - c) obtaining a database phenotype of the at least one database genetic sequence; and
- d) determining the phenotype of the <u>Human Immunodeficiency Virus</u> biological sample from the database phenotype.
- 29. (Currently amended) The method of claim 28, wherein the at least one database genetic sequence is at least about 60% identical to the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.
- 30. (Currently amended) The method of claim 29, wherein the at least one database genetic sequence is at least about 70% identical to the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.
- 31. (Currently amended) The method of claim 30, wherein the at least one database genetic sequence is at least about 80% identical to the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.
- 32. (Currently amended) The method of claim 31, wherein the at least one database genetic sequence is at least about 90% identical to the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample.
 - 33.-38. Canceled

DOCKET NO.: TIBO-0029 **Application No.:** 09/836,477

Office Action Dated: August 13, 2004

39. (Currently amended) A computer program for determining a phenotype of a retrovirus, wherein the retrovirus is the Human Immunodeficiency Virus, wherein the program is comprised on a computer readable medium biological sample, comprising:

- a) receiving a genetic sequence from [a] the Human Immunodeficiency Virus from the patient;
- b) identifying a mutation pattern of the genetic sequence of the <u>Human</u>

 <u>Immunodeficiency Virus</u> biological sample, wherein said mutation pattern comprises at least one mutation that correlates to resistance to at least one therapy;
- c) searching a relational genotype/phenotype database for at least one database mutation pattern similar to said mutation pattern of the genetic sequence of the <u>Human Immunodeficiency Virus</u> biological sample;
- d) obtaining at least one database phenotype of the at least one database mutation pattern from the relational genotype/phenotype database;
- e) determining the at least one phenotype of <u>Human Immunodeficiency Virus</u> biological sample from the at least one database phenotype; and
 - f) providing the phenotype of the <u>Human Immunodeficiency Virus</u> biological-sample.
- 40. (Original) The computer program of claim 39, wherein a series of phenotypes is obtained by repeating steps b) through e) for a group of therapies.
- 41. (Currently amended) The computer program of claim 40, wherein the phenotype of the <u>Human Immunodeficiency Virus</u> biological sample is provided in a report.
 - 42. Canceled